Page 2

The following <u>Listing of the Claims</u> will replace all prior versions and all prior listings of the claims in the present application:

<u>Listing of The Claims</u>:

- 1. (Withdrawn) One or more isolated nucleic acid sequences selected from the group consisting of those sequences identified in Figure 6A which correspond to genes 1-5807 identified in Figure 6 and/or those sequences identified in Figure 13.
- 2. (Withdrawn) A vector comprising an isolated nucleic acid sequence of claim 1.
- 3. (Withdrawn) A host cell comprising the vector of claim 2.
- 4. (Withdrawn) A composition comprising one or more chondrocyte enriched or chondrocyte-specific nucleic acid sequences isolated from one or more sources selected from the group consisting of (a) fetus, (b) normal, (c) mild osteoarthritic, (d) moderate osteoarthritic, (e) marked osteoarthritic and (f) severe osteoarthritic cartilage samples.
- 5. (Withdrawn) A composition comprising one or more nucleic acid sequences selected from the group consisting of sequences identified in Figure 6B whose sequences are disclosed in Figure 14.
- 6. (Withdrawn) A composition comprising one or more nucleic acid sequences selected from the group consisting of sequences identified in Figure 6C whose sequences are disclosed in Figure 14.
- 7. (Withdrawn) A composition comprising one or more nucleic acid sequences selected from the group consisting of sequences identified in Figure 6D whose sequences are disclosed in Figure 14.
- 8. (Withdrawn) A composition comprising one or more nucleic acid sequences selected from the group consisting of those sequences identified in Figure 6E whose sequences are disclosed in Figure 14.
- 9. (Withdrawn) A composition comprising one or more nucleic acid sequences selected from the group consisting of those sequences identified in Figure 6B, 6C, 6D, and 6E whose sequences are disclosed in Figure 14.

Page 3

10. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with mild osteoarthritis relative to cartilage from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue obtained less than 14 hours post-mortem.

- 11. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with severe osteoarthritis relative to cartilage derived from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue obtained less than 14 hours post-mortem.
- 12. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with marked osteoarthritis relative to cartilage derived from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue obtained less than 14 hours post-mortem.
- 13. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with moderate osteoarthritis relative to cartilage derived from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue obtained less than 14 hours post-mortem.
- 14. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with mild osteoarthritis relative to cartilage isolated from a fetus.
- 15. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with moderate osteoarthritis relative to cartilage isolated from a fetus.

Page 4

16. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with marked osteoarthritis relative to cartilage isolated from a fetus.

- 17. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from a patient diagnosed with severe osteoarthritis relative to cartilage isolated from a fetus.
- 18. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage from normal individual relative to cartilage isolated from a fetus.
- 19. (Withdrawn) A composition comprising one or more nucleic acid sequences wherein at least one of said nucleic acid sequences is differentially expressed in cartilage isolated from any two or more of the following sources: (a) fetus (b) patient with mild osteoarthritis, (c) patient with moderate osteoarthritis, (d) patient with marked osteoarthritis, (e) patient with severe osteoarthritis and (f) cartilage isolated from a normal individual isolated from cartilage tissue obtained less than 14 hours post-mortem.
- 20. (Withdrawn) A composition comprising one or more nucleic acid sequences identified in Figure 9 and/or sequences identified in Figure 6A which correspond to the genes disclosed in Figure 9.
- 21. (Withdrawn) A composition comprising one or more nucleic acid sequences identified in Figure 11 and/or sequences identified in Figure 6A which correspond to the genes disclosed in Figure 11.
- 22. (Withdrawn) A composition comprising one or more nucleic acid sequences identified in Figure 6A which correspond to the genes disclosed in Figure 15 and Figure 16.
- 23. (Withdrawn) A composition comprising one or more nucleic acids sequences identified in Figure 6A which correspond to the genes disclosed in Figure 6.
- 24. (Withdrawn) A composition comprising one or more nucleic acid sequences comprising one or more of the sequences disclosed in Figure 13.

Page 5

25. (Withdrawn) An array comprising:

a plurality of nucleic acid members and a solid substrate, wherein at least one member is differentially expressed in cartilage from a patient diagnosed with mild osteoarthritis as compared to cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

26. (Withdrawn) An array comprising:

a plurality of nucleic acid members and a solid substrate, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with severe osteoarthritis as compared to cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

27. (Withdrawn) An array comprising:

a plurality of nucleic acid members and a solid substrate, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with moderate osteoarthritis as compared to cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

28. (Withdrawn) An array comprising:

a plurality of nucleic acid members and a solid substrate, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with marked osteoarthritis as compared to cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

29. (Withdrawn) An array comprising:

a plurality of nucleic acid members and a solid support, wherein at least one member is differentially expressed in cartilage isolated from a fetus as compared to cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

30. (Withdrawn) An array comprising:

Page 6

a plurality of nucleic acid members and a solid substrate, wherein at least one member is differentially expressed in cartilage isolated from any two or more of the following sources: (a) fetus, (b) patient with mild osteoarthritis (c) patient with moderate osteoarthritis (d) patient with marked osteoarthritis (e) severe osteoarthritis, and (f) cartilage from a normal individual, and wherein each nucleic acid member has a unique position on said array and is stably associated with said solid substrate.

- 31. (Withdrawn) The array of claim 25, 26, 27, 28 29 or 30 wherein said normal individual is living.
- 32. (Withdrawn) The array of claim 25, 26, 27, 28, 29 or 30 wherein said cartilage isolated from said normal individual is isolated from cartilage tissue less than 14 hours postmortem.
- 33. (Withdrawn) The array of claim 25, 26, 27, 28, 29, 30 wherein each nucleic acid member is at least 50 nucleotides.
- 34. (Withdrawn) The array of claim 25, 26, 27, 28, 29, 30 wherein said array comprises from 10 to 20,000 positions.
- 35. (Withdrawn) The array of claim 25, 26, 27, 28, 29, 30 further including negative and positive control sequences and RNA quality control sequences selected from the group consisting of: cDNA sequences encoded by housekeeping genes, plant gene sequences, bacterial sequences, PCR products and vector sequences.
- 36. (Cancelled) A method of diagnosing mild osteoarthritis in a patient, comprising: hybridizing a nucleic acid sample corresponding to RNA to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with mild osteoarthritis, as compared to cartilage isolated from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue less than 14 hours post-mortem, and wherein each nucleic acid member has a unique position and is stably associated with the solid substrate, and wherein hybridization of said nucleic acid sample to one or more said differentially expressed nucleic acid members is indicative of mild osteoarthritis.

Page 7

- 37. (Cancelled A method of diagnosing moderate osteoarthritis in a patient comprising: hybridizing a nucleic acid sample corresponding to RNA to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with moderate osteoarthritis, as compared to cartilage isolated from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue less than 14 hours post-mortem, and wherein each nucleic acid member has a unique position and is stably associated with said solid substrate, and wherein hybridization of said nucleic acid sample to one or more said differentially expressed nucleic acid members is indicative of moderate osteoarthritis.
- 38. (Cancelled) A method of diagnosing marked osteoarthritis in a patient comprising: hybridizing a nucleic acid sample corresponding to RNA to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with marked osteoarthritis, as compared to cartilage isolated from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue less than 14 hours post-mortem, wherein each nucleic acid member has a unique position and is stably associated with said solid substrate, and wherein hybridization of said nucleic acid sample to one or more said differentially expressed nucleic acid members is indicative of marked osteoarthritis.
- 39. (Cancelled) A method of diagnosing severe osteoarthritis in a patient comprising: hybridizing a nucleic acid sample corresponding to RNA to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member is differentially expressed in cartilage isolated from a patient diagnosed with severe osteoarthritis, as compared to cartilage isolated from a normal individual, wherein cartilage isolated from said normal individual is isolated from cartilage tissue less than 14 hours post-mortem, wherein each nucleic acid member has a unique position and is stably associated with the solid substrate, and wherein hybridization of said nucleic acid sample to one or more said differentially expressed nucleic acid members is indicative of severe osteoarthritis.

Page 8

40. (Cancelled) The method of claim 36, 37, 38 or 39 further comprising the step of isolating RNA from said patient.

- 41. (Cancelled) The method of claim 40 further comprising the step of isolating RNA from a cartilage sample.
- 42. (Cancelled) The method of claim 40 further comprising the step of isolating RNA from a blood sample.
- 43. (Cancelled) The method of claim 40 further comprising the step of isolating RNA from a synovial fluid sample.
- 44. (Cancelled) The method of claim 36, 37, 38, or 39 further comprising the step of preparing a nucleic acid sample corresponding to the said RNA.
- 45. (Withdrawn) A method of identifying an agent that increases or decreases the expression of a nucleic acid sequence that is differentially expressed in a chondrocyte derived from any two or more of the following chondrocyte disease or developmental stages: fetal, mild osteoarthritic, moderate osteoarthritic, marked osteoarthritic and severe osteoarthritic, comprising:

incubating a chondrocyte derived from a normal individual with a candidate agent, wherein said chondrocyte is isolated from a cartilage sample obtained from said normal individual less than 14 hours post-mortem; isolating RNA from said chondrocyte; and hybridizing a probe to said RNA, said probe corresponding to a nucleic acid sequence which is differentially expressed in a chondrocyte derived from any two or more sources selected from the group consisting of: fetal, normal, mild osteoarthritic, moderate osteoarthritic, marked osteoarthritic and severe osteoarthritic, wherein differential hybridization of said probe to said RNA from said normal individual relative to RNA from one or more of fetal, mild osteoarthritic, marked osteoarthritis moderate osteoarthritis or severe osteoarthritic samples is indicative of the level of expression of RNA corresponding to a differentially expressed chondrocyte-specific nucleic acid sequence, and wherein, as a result of said incubation step in the presence of said candidate agent, a change in the level of expression of said nucleic acid sequence is indicative of an agent that increases or decreases the expression of said chondrocyte specific nucleic acid sequence.

Page 9

46. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,

- a) isolating chondrocytes from a cartilage sample derived from one or more normal individuals, wherein said cartilage sample is obtained less than 14 hours post-mortem;
- b) isolating mRNA from said chondrocytes;
- c) synthesizing cDNA from said mRNA; and
- d) ligating said cDNA into a vector.
- 47. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,
 - a) isolating chondrocytes from a cartilage sample derived from one or more living normal individuals;
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and
 - d) ligating said cDNA into a vector.
- 48. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,
 - a) isolating chondrocytes from a cartilage sample derived from one or more patients diagnosed with mild osteoarthritis
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and
 - d) ligating said cDNA into a vector.
- 49. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,
 - a) isolating chondrocytes from a cartilage sample derived from one or more patients diagnosed with moderate osteoarthritis
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and
 - e) ligating said cDNA into a vector.
- 50. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,
 - a) isolating chondrocytes from a cartilage sample derived from one or more patients diagnosed with marked osteoarthritis
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and

Page 10

- d) ligating said cDNA into a vector.
- 51. (Withdrawn) A method of preparing a chondrocyte cDNA library comprising,
 - a) isolating chondrocytes from a cartilage sample derived from one or more patients diagnosed with severe osteoarthritis
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and
 - f) ligating said cDNA into a vector.
- 52. (Withdrawn) A method of preparing a chondroctye cDNA library comprising,
 - a) isolating chondrocytes from one or more fetuses;
 - b) isolating mRNA from said chondrocytes;
 - c) synthesizing cDNA from said mRNA; and
 - d) ligating said cDNA into a vector.
- 53. (Withdrawn) A method of making an array comprising a plurality of nucleic acid members selected from those sequences identified in Figure 14 on a solid support, said support comprising a surface with a plurality of pre-selected unique regions, said method comprising:

spotting each nucleic acid member individually onto a unique pre-selected region and stably attaching each nucleic acid member to said solid support.

- 54. (Withdrawn) The method of claim 53, wherein at least one nucleic acid member is differentially expressed in cartilage isolated from two or more sources selected from the group consisting of: (a) a fetus or a patient diagnosed with (b) mild, (c) moderate, (d) marked, (e) severe osteoarthritis, and (f) cartilage isolated from a normal individual.
- 55. (Withdrawn) The method of claim 54, wherein the cartilage is isolated from one or more normal individuals is isolated from cartilage tissue less than 14 hours post-mortem.
- 56. (Withdrawn) A method of claim 54 wherein the cartilage is isolated from one or more living normal individuals.
- 57. (Withdrawn) A kit comprising an array of claim 25, 26, 27, 28, 29, 30 and packaging means therefore.

Page 11

- 58. (currently amended) A method of diagnosing human osteoarthritis (OA) in an individual, comprising determining the level of two or more-RNA transcripts which correpond to two or more genes respectively, selected from Figure 6-in a cartilage sample from an individual suspected of having or being afflicted with OA, wherein a difference in expression of said wherein the RNA transcripts are expressed from two or more RNA transcripts in said individual compared to the expression of said two or more RNA transcripts in an individual genes which are differentially expressed in OA cartilage as compared with normal cartilage as identified in Table 6, wherein said two or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing said level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having OA, wherein differential expression of said two or more RNA transcripts is indicative of the disease OA.
- 59. (currently amended) The method of claim 58, comprising determining the level of a plurality of RNA transcripts which correspond to where said RNA transcripts are expressed from three or more genes differentially expressed in OA cartilage as compared with normal cartilage as identified in Table 6, wherein differential expression of said three or more RNA transcripts is indicative of the disease OA, selected from Figure 6 in a sample from an individual suspected of having or being afflicted with OA, wherein a difference in expression of said plurality of RNA transcripts in said individual compared to the expression of said plurality of RNA transcripts in a normal individual is indicative of OA, and wherein said three or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1);

Page 12

MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1

(PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP).

- (currently amended) A method of diagnosing mild human osteoarthritis (OA) in an 60. individual, comprising determining the level of two or more RNA transcripts corresponding to two or more genes respectively, selected from Figure 6-in a cartilage sample from an individual suspected of having or being afflicted with mild OA, wherein a difference in expression of said wherein the RNA transcripts are expressed from two or more RNA transcripts in said individual compared to the expression of said two or more RNA transcripts in an individual genes which are differentially expressed in mild OA cartilage as compared with non-mild OA cartilage as identified in Table 6, wherein said two or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing the level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having mild OA, wherein differential expression of said two or more RNA transcripts is indicative of the disease mild OA.
- 61. (currently amended) The method of claim 60, eomprising determining the level of a plurality of RNA transcripts corresponding to where said RNA transcripts are expressed from three or more genes differentially expressed in mild OA cartilage as compared with non mild OA cartilage as identified in Table 6, wherein differential expression of said three or more RNA transcripts is indicative of the disease mild OA, and wherein said three or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP) selected from Figure 6 in

Page 13

a sample from an individual suspected of having or being afflicted with OA, wherein a difference in expression of said plurality of RNA transcripts in said individual compared to the expression of said plurality of RNA transcripts in a normal individual is indicative of OA.

- 62. (cancelled)
- 63. (cancelled)
- 64. (cancelled)
- 65. (cancelled)
- (currently amended) A method of diagnosing severe human osteoarthritis (OA) in an 66. individual, comprising determining the level of two or more RNA transcripts corresponding to two or more genes respectively, selected from Figure 6 in a cartilage sample from an individual suspected of having or being afflicted with severe OA, wherein a difference in expression of said wherein the RNA transcripts are expressed from two or more RNA transcripts in said individual compared to the expression of said two or more RNA transcripts in an individual genes which are differentially expressed in severe OA cartilage as compared with non severe OA cartilage as identified in Table 6, wherein said two or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); Bcell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing the level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having severe OA, wherein differential expression of said two or more RNA transcripts is indicative of the disease severe OA.
- 67. (currently amended) The method of claim 66, comprising determining the level of a plurality of RNA transcripts corresponding to where said RNA transcripts are expressed from three or more genes differentially expressed in severe OA cartilage as compared with non severe OA cartilage as identified in Table 6, wherein differential expression of said three or more RNA transcripts is indicative of the disease severe OA, and wherein

Page 14

said three or more genes are selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP). selected from Figure 6 in a sample from an individual suspected of having or being afflicted with severe OA, wherein a difference in expression of said plurality of RNA transcripts in said individual.

- 68. (currently amended) The method of any one of claims 58, 60, [62, 64] or 66, further comprising the step of isolating RNA from said patient cartilage sample.
- 69. (cancelled) The method of claim 68, further comprising the step of isolating RNA from a blood sample.
- 70. (cancelled) The method of claim 68, further comprising the step of isolating RNA from a synovial fluid sample.
- 71. (cancelled) The method of claim 68, further comprising the step of isolating RNA from said cartilage sample.
- 72. (cancelled) The method of claim 68, wherein said cartilage sample comprises cartilage isolated from cartilage tissue less than 14 hours post-mortem.
- 73.. (currently amended) The method of any one of claims 58, 60, [62, 64] or 66, wherein said the step of determining the level of said RNA transcripts comprises hybridizing a nucleic acid sample comprising or corresponding to said RNA transcripts corresponding to genes selected from Figure 6 from an individual suspected of having or being afflicted with OA, to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least two of said nucleic acid members are differentially expressed in cartilage isolated from a patient diagnosed with osteoarthritis as compared to cartilage isolated from a normal individual, wherein each nucleic acid member has a unique position and is stably associated with the solid substrate, and wherein hybridization of said nucleic acid sample to said differentially expressed nucleic acid members is indicative of osteoarthritis.

Page 15

- 74. (new) A method of diagnosing human osteoarthritis (OA), comprising determining the level of RNA transcripts in a cartilage sample from an individual suspected of having or being afflicted with OA, where the RNA transcripts are expressed from a plurality of the genes identified in Table 6 as being differentially expressed in OA cartilage as compared with normal cartilage, wherein said plurality of genes are selected from Beta 2

 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing the level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having OA, wherein the expression pattern of said plurality of RNA transcripts in said individual suspected of having OA is indicative of the disease OA.
 - 75. (new) A method of diagnosing mild human osteoarthritis (OA), comprising determining the level of RNA transcripts in a cartilage sample from an individual suspected of having or being afflicted with mild OA, where the RNA transcripts are expressed from a plurality of the genes identified in Table 6 as being differentially expressed in mild OA cartilage as compared with non-mild OA cartilage, wherein said plurality of genes are selected from Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing the level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having mild OA, wherein the expression pattern of said plurality of RNA transcripts in said individual suspected of having mild OA is indicative of the disease mild OA.
- 76. (new) A method of diagnosing severe human osteoarthritis (OA), comprising determining the level of RNA transcripts in a cartilage sample from an individual suspected of having

Page 16

or being afflicted with severe OA, where the RNA transcripts are expressed from a plurality of the genes identified in Table 6 as being differentially expressed in OA cartilage as compared with non-severe OA cartilage, wherein said plurality of genes are selected from Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and comparing the level of said RNA transcripts with the level of expression of said RNA transcripts in a cartilage sample from one or more individuals not having severe OA, wherein the expression pattern of said plurality of RNA transcripts in said individual suspected of having severe OA is indicative of the disease severe OA.

- 77. (new) A method for diagnosing human osteoarthritis (OA) in a patient comprising hybridizing a nucleic acid sample corresponding to RNA obtained from said patient to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member hybridizes to a nucleic acid target a gene which is differentially expressed in cartilage isolated from a patient diagnosed with OA as compared to cartilage isolated from a normal individual, as identified in Table 6, and wherein said gene is further selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and wherein differential hybridization of said nucleic acid target to one or more nucleic acid members is indicative of osteoarthritis.
- 78. (new) A method for diagnosing human mild osteoarthritis (OA) in a patient comprising hybridizing a nucleic acid sample corresponding to RNA obtained from said patient to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member hybridizes to a nucleic acid target from a gene which is differentially expressed in cartilage isolated from a patient diagnosed with mild OA, as compared to

Page 17

cartilage isolated from an individual without mild OA as identified in Table 6, and wherein said gene is further selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and wherein differential hybridization of said nucleic acid target to one or more nucleic acid members is indicative of mild osteoarthritis.

79. (new) A method for diagnosing severe osteoarthritis (OA) in a patient comprising hybridizing a nucleic acid sample corresponding to RNA obtained from said patient to an array comprising a solid substrate and a plurality of nucleic acid members, wherein at least one member hybridizes to a nucleic acid target from a gene which is differentially expressed in cartilage isolated from a patient diagnosed with severe OA, as compared to cartilage isolated from an individual without severe OA as identified in Table 6, and wherein said gene is further selected from the group consisting of Beta 2 Microglobulin (B2M); Tumour Necrosis Factor Alpha-induced Protein (TNFAIP6); B-cell CLL/lymphoma 6 (zinc finger protein 51) (BCL6); Cyclin C (CCNC); Interleukin 13 receptor alpha 1 (IL13RA1); Laminin, gamma 1 (LAMC1); MAFB/Kreisler basic region/leucine zipper transcription factor (MAFB); Period 1 (PER1); Calmodulin 1 (CALM1); Translationally Controlled Tumour Protein (TCTP), and wherein differential hybridization of said nucleic acid target to one or more nucleic acid members is indicative of severe osteoarthritis.